3/25/05

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* * * * * * * * *
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NEWS
NEWS
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      3
         SEP 01
                 New pricing for the Save Answers for SciFinder Wizard within
                 STN Express with Discover!
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         OCT 28
                 KOREAPAT now available on STN
         NOV 30 PHAR reloaded with additional data
NEWS
         DEC 01
NEWS
                 LISA now available on STN
         DEC 09
NEWS
                 12 databases to be removed from STN on December 31, 2004
      7
      8 DEC 15
NEWS
                 MEDLINE update schedule for December 2004
         DEC 17
NEWS
                 ELCOM reloaded; updating to resume; current-awareness
                  alerts (SDIs) affected
      10 DEC 17
NEWS
                 COMPUAB reloaded; updating to resume; current-awareness
                  alerts (SDIs) affected
NEWS
      11 DEC 17
                 SOLIDSTATE reloaded; updating to resume; current-awareness
                  alerts (SDIs) affected
      12 DEC 17
NEWS
                 CERAB reloaded; updating to resume; current-awareness
                  alerts (SDIs) affected
NEWS
      13 DEC 17
                 THREE NEW FIELDS ADDED TO IFIPAT/IFIUDB/IFICDB
      14 DEC 30
NEWS
                 EPFULL: New patent full text database to be available on STN
      15 DEC 30
NEWS
                 CAPLUS - PATENT COVERAGE EXPANDED
NEWS
      16 JAN 03
                 No connect-hour charges in EPFULL during January and
                  February 2005
NEWS
      17 FEB 25
                 CA/CAPLUS - Russian Agency for Patents and Trademarks
                  (ROSPATENT) added to list of core patent offices covered
NEWS
      18 FEB 10
                 STN Patent Forums to be held in March 2005
NEWS
      19 FEB 16
                 STN User Update to be held in conjunction with the 229th ACS
                 National Meeting on March 13, 2005
NEWS
      20 FEB 28
                 PATDPAFULL - New display fields provide for legal status
                 data from INPADOC
NEWS
      21 FEB 28
                 BABS - Current-awareness alerts (SDIs) available
NEWS
      22 FEB 28
                 MEDLINE/LMEDLINE reloaded
NEWS
      23 MAR 02
                 GBFULL: New full-text patent database on STN
                 REGISTRY/ZREGISTRY - Sequence annotations enhanced
NEWS
      24 MAR 03
NEWS
      25 MAR 03
                 MEDLINE file segment of TOXCENTER reloaded
      26 MAR 22
NEWS
                 KOREAPAT now updated monthly; patent information enhanced
      27 MAR 22
                 Original IDE display format returns to REGISTRY/ZREGISTRY
NEWS
NEWS
      28 MAR 22
                 PATDPASPC - New patent database available
NEWS
                 REGISTRY/ZREGISTRY enhanced with experimental property tags
      29 MAR 22
NEWS EXPRESS
              JANUARY 10 CURRENT WINDOWS VERSION IS V7.01a, CURRENT
              MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
              AND CURRENT DISCOVER FILE IS DATED 10 JANUARY 2005
NEWS HOURS
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              CAS World Wide Web Site (general information)
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FILE 'HOME' ENTERED AT 12:12:09 ON 25 MAR 2005

=> file registry COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 12:12:34 ON 25 MAR 2005 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

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Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 24 MAR 2005 HIGHEST RN 847222-24-6 DICTIONARY FILE UPDATES: 24 MAR 2005 HIGHEST RN 847222-24-6

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 18, 2005

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

```
=> e tibolone/cn
E1
            1
                  TIBO-R 86183/CN
                  TIBOLON LIVIELLA/CN
E2
            1
            1 --> TIBOLONE/CN
E3
                 TIBON/CN
Ε4
            1
E5
                 TIBONE/CN
            1
                 TIBOR 2.5-2.5/CN
Ε6
            1
                TIBOR 3-0.2/CN
E7
            1
           1
                TIBOR 3-1/CN
E8
           1
                TIBOR 33/CN
E9
           1
                TIBOR 5-0.2/CN
E10
                TIBOR 5-0.6/CN
E11
           1
           1
                TIBOR 5-1/CN
E12
```

=> s e2-e3

E1

1

8

1 "TIBOLON LIVIELLA"/CN

1 TIBOLONE/CN

L1 2 ("TIBOLON LIVIELLA"/CN OR TIBOLONE/CN)

=> file caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL

ENTRY SESSION

9.84

FULL ESTIMATED COST 9.63

FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 25 Mar 2005 VOL 142 ISS 14 FILE LAST UPDATED: 24 Mar 2005 (20050324/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

```
=> s 11
           386 L1
L2
=> e tibolone/bi
E1
                   TIBOLIUM/BI
           2
E2
             3
                  TIBOLON/BI
           358 --> TIBOLONE/BI
E3
                 TIBOLONES/BI
E4
           1
E5
            14
                  TIBON/BI
E6
                 TIBONCHINA/BI
            1
E7
             5
                 TIBOND/BI
E8
             1
                 TIBONDS/BI
                 TIBONE/BI
E9
            1
                 TIBONI/BI
E10
            1
E11
             1
                  TIBONUCLEOPROTEIN/BI
E12
             4
                   TIBOOBURRA/BI
=> s e2-e4
             3 TIBOLON/BI
           358 TIBOLONE/BI
            1 TIBOLONES/BI
           358 TIBOLONE/BI
                 ((TIBOLONE OR TIBOLONES)/BI)
             1 TIBOLONES/BI
           359 (TIBOLON/BI OR TIBOLONE/BI OR TIBOLONES/BI)
1.3
=> e tibolone/ct
E#
    FREQUENCY
                  AT
                         TERM
                  __
```

TIBICEN WINNEMANNA/CT

```
0
E2
                       TIBIELLA/CT
            0
                    --> TIBOLONE/CT
E3
E4
             0
                  1 TIBOUCHIA/CT
E5
             0
                 2
                       TIBOUCHIA BOURGAENA/CT
            1 13
                       TIBOUCHINA/CT
E6
                 2
E7
            0
                       TIBOUCHINA ALIENA/CT
E8
            0
                       TIBOUCHINA GRANDIFLORA/CT
            4
                 8
                       TIBOUCHINA GRANULOSA/CT
E9
            1
                16
                       TIBOUCHINA LONGIFOLIA/CT
E10
            2
                 8
                       TIBOUCHINA MULTIFLORA/CT
E11
                 8
E12
           12
                        TIBOUCHINA PULCHRA/CT
=> s 12 or 13
          430 L2 OR L3
L4
=> s starch? or (potato starch?) or (corn starch?) or (amylomaize starch?) or
amylum? or keoflo? or (wheat starch?) or (modified starch?) or (agglomerated
starch?) or (granulated starch?)
        147702 STARCH?
         53947 POTATO
            35 POTATOS
         23309 POTATOES
         61080 POTATO
                 (POTATO OR POTATOS OR POTATOES)
        147702 STARCH?
          8524 POTATO STARCH?
                 (POTATO(W)STARCH?)
        112878 CORN
           365 CORNS
        113011 CORN
                 (CORN OR CORNS)
        147702 STARCH?
         10943 CORN STARCH?
                 (CORN(W)STARCH?)
           192 AMYLOMAIZE
             5 AMYLOMAIZES
           192 AMYLOMAIZE
                 (AMYLOMAIZE OR AMYLOMAIZES)
        147702 STARCH?
           83 AMYLOMAIZE STARCH?
                 (AMYLOMAIZE (W) STARCH?)
           114 AMYLUM?
             1 KEOFLO?
        116352 WHEAT
          2802 WHEATS
        116449 WHEAT
                 (WHEAT OR WHEATS)
        147702 STARCH?
          3221 WHEAT STARCH?
                 (WHEAT (W) STARCH?)
        511063 MODIFIED
             5 MODIFIEDS
        511066 MODIFIED
                 (MODIFIED OR MODIFIEDS)
        147702 STARCH?
          2982 MODIFIED STARCH?
                 (MODIFIED (W) STARCH?)
          7355 AGGLOMERATED
        147702 STARCH?
             5 AGGLOMERATED STARCH?
                 (AGGLOMERATED (W) STARCH?)
         30570 GRANULATED
        147702 STARCH?
           15 GRANULATED STARCH?
```

```
(GRANULATED (W) STARCH?)
L5
        147787 STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
               STARCH?) OR AMYLUM? OR KEOFLO? OR (WHEAT STARCH?) OR (MODIFIED
               STARCH?) OR (AGGLOMERATED STARCH?) OR (GRANULATED STARCH?)
=> s stabilizer? or (stabilizing (W) (agent? or component? or compound?)) or
(antioxida?) or (chelat?)
        105727 STABILIZER?
         67792 STABILIZING
             2 STABILIZINGS
         67794 STABILIZING
                 (STABILIZING OR STABILIZINGS)
       1461153 AGENT?
       1121008 COMPONENT?
        895362 COMPOUND?
       1046326 COMPD
       1610396 COMPDS
       2282540 COMPD
                 (COMPD OR COMPDS)
       2681023 COMPOUND?
                 (COMPOUND? OR COMPD)
         16402 STABILIZING (W) (AGENT? OR COMPONENT? OR COMPOUND?)
        125806 ANTIOXIDA?
          2529 ANTIOXIDN
             4 ANTIOXIDNS
          2533 ANTIOXIDN
                 (ANTIOXIDN OR ANTIOXIDNS)
        126860 ANTIOXIDA?
                 (ANTIOXIDA? OR ANTIOXIDN)
        124446 CHELAT?
L6
        348947 STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMPOUN
               D?)) OR (ANTIOXIDA?) OR (CHELAT?)
=> s (ascorbyl palmitate?) or "6-0-palmitoylascorbic acid" or
"ascorbate-6-palmitate" or "vitamin C palmitate" or "ascorbic-6-0-palmitate" or
(ascorbyl stearate?) or (sodium ascorbate?) or (ascorbic acid?) or hybrin? or
"L-ascorbic acid"
          2114 ASCORBYL
         23179 PALMITATE?
          1124 ASCORBYL PALMITATE?
                 (ASCORBYL (W) PALMITATE?)
       3538616 "6"
       1444243 "O"
            59 "PALMITOYLASCORBIC"
       3954637 "ACID"
       1469590 "ACIDS"
       4430938 "ACID"
                 ("ACID" OR "ACIDS")
            11 "6-O-PALMITOYLASCORBIC ACID"
                 ("6"(W)"O"(W)"PALMITOYLASCORBIC"(W)"ACID")
         32312 "ASCORBATE"
           592 "ASCORBATES"
         32530 "ASCORBATE"
                 ("ASCORBATE" OR "ASCORBATES")
       3538616 "6"
         22899 "PALMITATE"
           479 "PALMITATES"
         23167 "PALMITATE"
                 ("PALMITATE" OR "PALMITATES")
             4 "ASCORBATE-6-PALMITATE"
                 ("ASCORBATE"(W)"6"(W)"PALMITATE")
        179642 "VITAMIN"
         50473 "VITAMINS"
        198641 "VITAMIN"
```

```
3283799 "C"
         22899 "PALMITATE"
           479 "PALMITATES"
         23167 "PALMITATE"
                 ("PALMITATE" OR "PALMITATES")
            49 "VITAMIN C PALMITATE"
                 ("VITAMIN"(W)"C"(W)"PALMITATE")
         77508 "ASCORBIC"
       3538616 "6"
       1444243 "O"
         22899 "PALMITATE"
           479 "PALMITATES"
         23167 "PALMITATE"
                 ("PALMITATE" OR "PALMITATES")
             0 "ASCORBIC-6-O-PALMITATE"
                 ("ASCORBIC"(W)"6"(W)"O"(W)"PALMITATE")
          2114 ASCORBYL
         67574 STEARATE?
           129 ASCORBYL STEARATE?
                  (ASCORBYL (W) STEARATE?)
        970377 SODIUM
            34 SODIUMS
        970386 SODIUM
                  (SODIUM OR SODIUMS)
         32534 ASCORBATE?
          1596 SODIUM ASCORBATE?
                 (SODIUM(W) ASCORBATE?)
         77508 ASCORBIC
       4628709 ACID?
         76833 ASCORBIC ACID?
                 (ASCORBIC (W) ACID?)
             1 HYBRIN?
       1395378 "L"
         77508 "ASCORBIC"
       3954637 "ACID"
       1469590 "ACIDS"
       4430938 "ACID"
                 ("ACID" OR "ACIDS")
         12316 "L-ASCORBIC ACID"
                 ("L"(W) "ASCORBIC"(W) "ACID")
L7
         78296 (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASCORB
               ATE-6-PALMITATE" OR "VITAMIN C PALMITATE" OR "ASCORBIC-6-O-PALMI
               TATE" OR (ASCORBYL STEARATE?) OR (SODIUM ASCORBATE?) OR (ASCORBI
               C ACID?) OR HYBRIN? OR "L-ASCORBIC ACID"
=> s lactose or disaccharide?
         47024 LACTOSE
            90 LACTOSES
         47032 LACTOSE
                 (LACTOSE OR LACTOSES)
         16651 DISACCHARIDE?
         61648 LACTOSE OR DISACCHARIDE?
L8
=> d his
     (FILE 'HOME' ENTERED AT 12:12:09 ON 25 MAR 2005)
     FILE 'REGISTRY' ENTERED AT 12:12:34 ON 25 MAR 2005
                E TIBOLONE/CN
L1
              2 S E2-E3
     FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005
L2
            386 S L1
```

("VITAMIN" OR "VITAMINS")

```
E TIBOLONE/BI
            359 S E2-E4
L3
                E TIBOLONE/CT
L4
             430 S L2 OR L3
L5
         147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L6
         348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
          78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L7
L8
          61648 S LACTOSE OR DISACCHARIDE?
=> s 14 and 15 and (16 or 17)
             5 L4 AND L5 AND (L6 OR L7)
=> d 19 1-5 ibib ed abs
     ANSWER 1 OF 5 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
                          2004:1036906 CAPLUS
DOCUMENT NUMBER:
                          142:11606
TITLE:
                          Immediate-release pharmaceutical dosage form
                          comprising a tibolone polymorph
INVENTOR(S):
                          De Haan, Pieter
PATENT ASSIGNEE(S):
                          Akzo Nobel N.V., Neth.
                          PCT Int. Appl., 32 pp.
SOURCE:
                          CODEN: PIXXD2
DOCUMENT TYPE:
                          Patent
LANGUAGE:
                          English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
     PATENT NO.
                          KIND
                                 DATE
                                            APPLICATION NO.
                                                                      DATE
                          ----
                                 _____
                                             ______
                                                                      _____
     WO 2004103378
                          A1 20041202 WO 2004-EP50852
                                                                      20040519
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
             SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
             SN, TD, TG
     CA 2473718
                                 20041123
                                            CA 2004-2473718
                           AΑ
                                                                      20040519
     EP 1499278
                                           EP 2004-737220
                          A1
                                 20050126
                                                                      20040519
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK,
                                                               A 20030523
W 20040519
PRIORITY APPLN. INFO.:
                                              EP 2003-101500
                                              WO 2004-EP50852
ED
     Entered STN: 03 Dec 2004
AB
     An immediate-release pharmaceutical dosage form comprises a
     tibolone polymorph and excipients, wherein the polymorph has a
     mean particle size as defined in the specification of below 22.8 \mu m in
     the dosage form. Thus, an immediate-release tablet formulation contained
     tibolone 2.5, potato starch 10,
     ascorbyl palmitate 0.2, Mg stearate 0.5, and lactose
     86.8 mg.
REFERENCE COUNT:
                          5
                                THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
                                RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 2 OF 5 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
                          2004:453011 CAPLUS
DOCUMENT NUMBER:
                          140:429053
TITLE:
                          Pharmaceutical tablets containing tibolone
                          and a coating
```

INVENTOR(S): De Haan, Pieter; Van Lare, Cornelis Elisabeth Johannus

PATENT ASSIGNEE(S): Akzo Nobel N.V., Neth. SOURCE: PCT Int. Appl., 20 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	PATENT NO.				KIND DATE			APPLICATION NO.					DATE					
WO	2004	0455	87		A1	_	2004	0603	1	WO 2	003-	EP50	- - 829		2	0031	113	
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		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
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		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	ΜZ,	NI,	NO,	
		NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,	ТJ,	
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		TR,	BF,	ВJ,	CF,	CG,	CI,	CM,				GW,		MR,	NE,	SN,	TD,	TG
PRIORITY	APP	LN.	INFO	.:						EP 2	002-	7992	0	7	A 2	0021	120	
									1	US 2	002-	4277	40P]	P 2	0021	120	

ED Entered STN: 04 Jun 2004

The present invention relates to a stabilized pharmaceutical tablet AΒ comprising an amount of from 0.1 to 10% by weight of tibolone provided with a coating, such as a film coating, a sugar coating, a sugar film coating or a wrap coating. The invention further relates to the use of a coating for stabilizing a pharmaceutical tablet comprising tibolone. Preferably, the coating is a sugar coating or a sugar film coating. Preferably, the sugar comprises sucrose. These tablets were shown to have a lower amount of OM38, i.e. an isomerization degradation product of tibolone, after storage as compared to uncoated tibolone tablets. For example, a tablet was formulated containing tibolone, basic granulates (starch/lactose), ascorbyl palmitate, and Mg stearate and a film-coated with a composition containing hydroxypropyl Me cellulose, polyethylene glycol, titania, and talc, further coated with a composition containing arabic gum, saccharose, Ca carbonate, talc, titania, polyethylene glycol, glucose syrup, glycerol, and china clay.

ANSWER 3 OF 5 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1998:708947 CAPLUS

DOCUMENT NUMBER:

129:335779

TITLE: Stabilized tibolone compositions

INVENTOR(S): Haan De, Pieter; Rovers, Adrianus Cornelis Petrus; Lambregts-De Hurk, Theodora Antonia Maria; Zwinkels,

Jocominus Antonius Maria; Morita, Ryoichi

Akzo Nobel N.V., Neth. PATENT ASSIGNEE(S): PCT Int. Appl., 21 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO			KIN	D	DATE			APPL	ICAT	ION	NO.		D	ATE	
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M	o, RU,	TJ,	TM												

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                                                                  19980420
    NZ 338028
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                               20011026
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    RU 2207133
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                               20030627
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    AT 281836
                        E
                               20041115
                                           AT 1998-928214
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    JP 10316573
                        A2
                               19981202
                                           JP 1998-112312
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PRIORITY APPLN. INFO.:
                                           EP 1997-201180
                                                               A 19970422
                                                              W 19980420
                                           WO 1998-EP2361
                                           US 1999-403139
                                                               Al 19991014
                                           US 2002-117899
                                                               A1 20020408
                                           US 2002-303494
                                                               B1 20021125
ED
     Entered STN: 09 Nov 1998
AΒ
     The invention pertains to a pharmaceutical dosage unit, such as a tablet
     or a capsule, comprising an effective amount of tibolone (0.1-10 %
    by weight) and a starch-containing pharmaceutically acceptable carrier
     (also denoted as basic granules), wherein the carrier contains 10-100% by
     weight of the starch. Thus, a more stable tibolone
     formulation is obtained, allowing dry storage and lower doses of active
     ingredient. Thus, basic granules contained tibolone 0.3, HPC
     1.95, corn starch 32.5, Mg stearate 0.325, and lactose
     to 65 mg.
REFERENCE COUNT:
                         5
                               THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
                              RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 4 OF 5 CAPLUS COPYRIGHT 2005 ACS on STN
                        1994:686615 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         121:286615
TITLE:
                         Pharmaceutical compositions containing pregnane
                        derivatives for the treatment of tumors.
INVENTOR(S):
                        Deckers, Godefridus Hermanus Johanna; Kloosterboer,
                        Helenius Jan
```

DOCUMENT TYPE: Patent
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT ASSIGNEE(S):

SOURCE:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 613687	A1	19940907	EP 1994-200523	19940302

Akazo Nobel N.V., Neth.

Eur. Pat. Appl., 9 pp.

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EP 613687
                          В1
                                19990602
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE
     CA 2116829
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                                            CA 1994-2116829
                          AA
                                                                   19940302
     CA 2116829
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     AU 9457542
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     JP 07002673
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                         A2
                                19950106
                                                                   19940304
     JP 3391539
                         B2
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     US 5512556
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                                19960430
                                            US 1994-358639
                                                                  19941219
                                                                A 19930305
PRIORITY APPLN. INFO.:
                                            EP 1993-200631
                                                                B1 19940307
                                            US 1994-206994
                         MARPAT 121:286615
OTHER SOURCE(S):
     Entered STN: 10 Dec 1994
ED
     Pharmaceutical compns. containing pregnane derivs. (Markush structure given)
AΒ
     are useful for prevention or treatment of tumors. A twice daily dose of 1
     mg/kg (7\alpha, 17\alpha)-17-hydroxy-7-methyl-19-nor-17-pregn-5(10)-en-20-
     yn-3-one (I) inhibited the tumor growth in rats up to .apprx.70%. A
     tablet contained I 2.5, starch 10, ascorbyl
     palmitate 0.2, Mg stearate 0.5, and lactose q.s. to 100 mg.
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ANSWER 5 OF 5 CAPLUS COPYRIGHT 2005 ACS on STN

1991:214521 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 114:214521

TITLE: Pharmaceutical composition containing the pure

crystalline forms of $(7\alpha, 17\alpha)-17$ -hydroxy-7-methyl-19-nor-17-pregn-5(10)-en-20-yn-3-one

Sas, G. A. J. M. T.; Van Doornum, E. M. INVENTOR(S):

PATENT ASSIGNEE(S): AKZO N. V., Neth.

Eur. Pat. Appl., 13 pp. SOURCE:

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 389035 EP 389035	A1 B1	19900926 19931215	EP 1990-200574	19900312
R: AT, BE, CH,	DE, DK	, ES, FR, C	GB, GR, IT, LI, NL, SE	
ZA 9001674	À	19901228	ZA 1990-1674	19900305
CA 2011452	С	19990525	CA 1990-2011452	19900305
AT 98490	E	19940115	AT 1990-200574	19900312
ES 2062292	Т3	19941216	ES 1990-200574	19900312
JP 03047195	A2	19910228	JP 1990-63921	19900314
JP 3139709	B2	20010305		
JP 2001139473	A2	20010522	JP 2000-316716	19900314
US 5037817	A	19910806	US 1990~494089	19900315
AU 9051394	A1	19900920	AU 1990-51394	19900316
AU 625083	B2	19920702		
FI 94865	В	19950731	FI 1990-1328	19900316
FI 94865	C	19951110		
KR 183401	B1	19990501	KR 1990-3574	19900317
PRIORITY APPLN. INFO.:			NL 1989-673	A 19890318
			EP 1990-200574 A	A 19900312
			JP 1990-63921	A3 19900314
ED Enternal COM. 21 May	1001			

ED Entered STN: 31 May 1991

AB Pure monoclinic or triclinic $(7\alpha, 17\alpha) - 17 - \text{hydroxy} - 7 - \text{methyl} - 19$ nor-17-pregn-5(10)-en-20-yn-3-one (I) are more suitable for pharmaceutical compns. than the polymorphous I. A solution of 1 kg I in 23 L acetone containing

6 mL pyridine, was poured, under N, in 25 L water, followed by cooling to 0-5°, to crystallize monoclinic I (crystalline structure and x-ray diffraction spectrum given). Tablets comprised monoclinic I 2.5, starch 10, ascorbyl palmitate 0.2, Mg stearate 0.5, and lactose to 100 mg/tablet. I forms are useful for treating menopausal disorders, for immunomodulation, and for treating osteoporosis (no data).

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=> d his
     (FILE 'HOME' ENTERED AT 12:12:09 ON 25 MAR 2005)
     FILE 'REGISTRY' ENTERED AT 12:12:34 ON 25 MAR 2005
                E TIBOLONE/CN
              2 S E2-E3
L1
     FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005
            386 S L1
L2
                E TIBOLONE/BI
            359 S E2-E4
L3
                E TIBOLONE/CT
            430 S L2 OR L3
L4
         147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L5
         348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
L6
          78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L7
          61648 S LACTOSE OR DISACCHARIDE?
L8
L9
              5 S L4 AND L5 AND (L6 OR L7)
=> s tibolone? and starch?
           358 TIBOLONE?
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L10
             3 TIBOLONE? AND STARCH?
=> s 110 not 19
L11
             0 L10 NOT L9
=> s 14 and 18
             7 L4 AND L8
L12
=> s 112 not 19
             2 L12 NOT L9
L13
=> d 113 1-2 ibib ed abs
L13 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
                         2004:182541 CAPLUS
DOCUMENT NUMBER:
                         140:229476
TITLE:
                         Treatment of dyspareunia with topically administered
                         nitroglycerin formulations
INVENTOR(S):
                         Place, Virgil A.; Wilson, Leland F.; Doherty, Paul C.;
                         Hanamoto, Mark S.; Spivack, Alfred P.; Gesundheit,
                         Neil; Bennett, Sean R.; Doherty, Jane K.
PATENT ASSIGNEE(S):
                         USA
                         U.S. Pat. Appl. Publ., 18 pp., Cont.-in-part of U.S.
SOURCE:
                         Ser. No. 905,458.
                         CODEN: USXXCO
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
                         6
PATENT INFORMATION:
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PATENT NO. KIND DATE APPLICATION NO. DATE

US 2004044080	A1	20040304	US	2003-407858	20030404
US 5877216	A	19990302	US	1997-959064	19971028
US 6306841	B1	20011023	US	2000-539484	20000330
US 2001051656	A1	20011213	US	2001-905458	20010713
US 6593313	B2	20030715			
PRIORITY APPLN. INFO.:			US	1997-959057	B2 19971028
			US	1997-959064	A2 19971028
			US	1998-181316	B1 19981027
			US	2000-539484	A1 20000330
			US	2001-905458	A2 20010713

ED Entered STN: 05 Mar 2004

AB Methods and formulations for treating dyspareunia are provided. A pharmaceutical composition formulated so as to contain a therapeutically effective amount of nitroglycerin is administered to the vagina or vulvar area of the individual undergoing treatment. Preferred formulations are immediate release formulations in which at least 80% of the nitroglycerin in the formulation is released therefrom within 4 h following administration. The formulations may contain one or more addnl. active agents, e.g., agents that are also useful to treat dyspareunia and/or potentiate the action of nitroglycerin. Such addnl. agents include vasoactive agents such as prostaglandins, phosphodiesterase inhibitors, androgens such as testosterone, estrogens such as estradiol, and selective modulators of estrogen and androgen receptors. A kit for a patient to use in the self-administration of the formulation is also provided.

L13 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2001:856713 CAPLUS

DOCUMENT NUMBER: 136:178102

TITLE: A pilot study of hormone replacement therapy with

tibolone in women with mastopathic breasts

AUTHOR(S): Egarter, Christian; Eppel, Wolfgang; Vogel, Sonja;

Wolf, Georg

CORPORATE SOURCE: Department of Obstetrics and Gynecology, University of

Vienna, Vienna, 1010, Austria Maturitas (2001), 40(2), 165-171 CODEN: MATUDK; ISSN: 0378-5122

PUBLISHER: Elsevier Science Ireland Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English ED Entered STN: 27 Nov 2001

SOURCE:

The authors' purpose was to investigate the effect of tibolone AΒ in a prospective randomized, double-blind pilot study in postmenopausal women with clin. palpable mastopathic changes. Twenty postmenopausal non-users of hormone replacement therapy undergoing mammog. were randomly allocated to receive either 2.5 mg tibolone (Livial, Organon) or placebo (2 mg lactose). Mammog. d. according to the Wolfe classification, severity of breast discomfort, parity, smoking habits, body weight and FSH and estradiol (E2) levels were documented at baseline and after 6 mo of treatment. There were no statistically significant differences between the two treatment groups in terms of parity, smoking, body weight and FSH and E2 levels. At 6 mo, four patients in the tibolone group showed reduced breast d., compared with one patient in the placebo group. Whereas no significant differences were found between the two treatment groups when the categories 'minimal reduction' and 'reduction' in mammog. d. were combined, combination of the categories 'no change' and 'minimal reduction' showed a significant difference. Also, women in the tibolone group showed a statistically significant improvement in breast discomfort. This pilot study showed decreases in breast d. as well as an attendant significant alleviation of breast discomfort in women with baseline mastopathic changes treated with tibolone for 6 mo, which may be due to the inhibitory effect of tibolone on the enzymes involved in the biosynthesis of estradiol demonstrated in previous trials and should be further evaluated in long-term studies.

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(FILE 'HOME' ENTERED AT 12:12:09 ON 25 MAR 2005)

FILE 'REGISTRY' ENTERED AT 12:12:34 ON 25 MAR 2005

E TIBOLONE/CN

2 S E2-E3 L1

FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005

L2 386 S L1

E TIBOLONE/BI

359 S E2-E4 L3

E TIBOLONE/CT

430 S L2 OR L3

L5 147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE

348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP L6 L7 78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC

61648 S LACTOSE OR DISACCHARIDE?

L8 L9 5 S L4 AND L5 AND (L6 OR L7)

3 S TIBOLONE? AND STARCH? L10

L11 0 S L10 NOT L9

L12 7 S L4 AND L8

2 S L12 NOT L9 L13

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CONNECT CHARGES	6.63	7.15
NETWORK CHARGES	1.02	1.14
SEARCH CHARGES	107.73	116.93
DISPLAY CHARGES	18.55	18.55
FULL ESTIMATED COST	133.93	143.77
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
	ENTRY	SESSION

-5.11 -5.11

IN FILE 'CAPLUS' AT 12:23:14 ON 25 MAR 2005

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1 HAAN PIETER/AU
1 HAAN POUL DE/AU
3 HAAN R E/AU
1 HAAN RENE JOHAN/AU
1 HAAN ROBERT EDGAR/AU
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E7

E8 E9

E10 E11

E12

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1 "HAAN PIETER"/AU

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VAN DEN IJSSEL JAN/AU

VAN DEN IJSSEL PAUL/AU
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                24 "MORITA R Y"/AU
                14 "MORITA RYOICHI"/AU
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L1
                   2 S E2-E3
       FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005
L2
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                     E TIBOLONE/BI
L3
                359 S E2-E4
                     E TIBOLONE/CT
L4
                430 S L2 OR L3
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E21

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147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
         348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
          78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L7
          61648 S LACTOSE OR DISACCHARIDE?
L8
L9
              5 S L4 AND L5 AND (L6 OR L7)
L10
              3 S TIBOLONE? AND STARCH?
L11
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L12
              7 S L4 AND L8
L13
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                E VAN DEN HURK T/AU
              1 S E4
L17
                E MORITA R/AU
             97 S E3-E6, E23
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                E ROVERS A/AU
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L19
                E ZWINKELS J/AU
L20
              3 S E3, E8
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L21
=> s 121 or 116 or 117 or 118 or 119 or 120
           198 L21 OR L16 OR L17 OR L18 OR L19 OR L20
L22
=> s 122 and tibolone?
           358 TIBOLONE?
L23
             4 L22 AND TIBOLONE?
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L23 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
                          2004:1036906 CAPLUS
DOCUMENT NUMBER:
                          142:11606
TITLE:
                          Immediate-release pharmaceutical dosage form
                          comprising a tibolone polymorph
                          De Haan, Pieter
INVENTOR(S):
PATENT ASSIGNEE(S):
                          Akzo Nobel N.V., Neth.
                          PCT Int. Appl., 32 pp.
SOURCE:
                          CODEN: PIXXD2
DOCUMENT TYPE:
                          Patent
LANGUAGE:
                          English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
     PATENT NO.
                          KIND
                                 DATE
                                              APPLICATION NO.
                                                                       DATE
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     WO 2004103378
                          A1
                                 20041202 WO 2004-EP50852
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SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,

SN, TD, TG

CA 2473718 AA 20041123 CA 2004-2473718 20040519 EP 1499278 A1 20050126 EP 2004-737220 20040519

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR

PRIORITY APPLN. INFO.: EP 2003-101500 A 20030523 WO 2004-EP50852 W 20040519

ED Entered STN: 03 Dec 2004

AB An immediate-release pharmaceutical dosage form comprises a tibolone polymorph and excipients, wherein the polymorph has a mean particle size as defined in the specification of below 22.8 µm in the dosage form. Thus, an immediate-release tablet formulation contained tibolone 2.5, potato starch 10, ascorbyl palmitate 0.2, Mg stearate 0.5, and lactose 86.8 mg.

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L23 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

2004:999673 CAPLUS

DOCUMENT NUMBER:

141:415963

TITLE:

Immediate release dosage form for hormone replacement

therapy

INVENTOR(S):

De Haan, Pieter

PATENT ASSIGNEE(S):

Neth.

SOURCE: U.S.

U.S. Pat. Appl. Publ., 10 pp., Division of U.S. Ser.

No. 723,412, abandoned.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ______ -----____ _____ US 2004229854 A1 20041118 US 2004-781493 20040218 PRIORITY APPLN. INFO.: US 2000-723412 B3 20001128

ED Entered STN: 19 Nov 2004

AB It has been recognized that **tibolone**, which is an agent in the treatment of menopausal complaints and osteoporosis, if formulated with a particle size in a defined range, displays an unexpectedly high bioavailability of one of its estrogenic metabolites, Org A.

L23 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

2004:453011 CAPLUS

DOCUMENT NUMBER:

140:429053

TITLE:

Pharmaceutical tablets containing tibolone

and a coating

INVENTOR(S):

De Haan, Pieter; Van Lare, Cornelis

Elisabeth Johannus

PATENT ASSIGNEE(S): SOURCE:

Akzo Nobel N.V., Neth. PCT Int. Appl., 20 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAI	'ENT	NO.			KIN	D	DATE		1	APPL	ICAT:	ION I	NO.		Di	ATE	
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WO	2004	0455	87		A1		2004	0603	1	WO 2	003-1	EP50	329		20	0031	113
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            TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                           EP 2002-79920
                                                             A 20021120
                                           US 2002-427740P
                                                              P 20021120
```

Entered STN: 04 Jun 2004 ED

The present invention relates to a stabilized pharmaceutical tablet AB comprising an amount of from 0.1 to 10% by weight of tibolone provided with a coating, such as a film coating, a sugar coating, a sugar film coating or a wrap coating. The invention further relates to the use of a coating for stabilizing a pharmaceutical tablet comprising tibolone. Preferably, the coating is a sugar coating or a sugar film coating. Preferably, the sugar comprises sucrose. These tablets were shown to have a lower amount of OM38, i.e. an isomerization degradation product of tibolone, after storage as compared to uncoated tibolone tablets. For example, a tablet was formulated containing tibolone, basic granulates (starch/lactose), ascorbyl palmitate, and Mg stearate and a film-coated with a composition containing hydroxypropyl Me

cellulose, polyethylene glycol, titania, and talc, further coated with a composition containing arabic gum, saccharose, Ca carbonate, talc, titania, polyethylene glycol, glucose syrup, glycerol, and china clay.

L23 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN

1998:708947 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

129:335779

TITLE:

Stabilized tibolone compositions

INVENTOR(S):

Haan De, Pieter; Rovers, Adrianus Cornelis Petrus; Lambregts-De Hurk, Theodora Antonia

Maria; Zwinkels, Jocominus Antonius Maria;

Morita, Ryoichi

PATENT ASSIGNEE(S):

Akzo Nobel N.V., Neth. PCT Int. Appl., 21 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE																	
WO	9847	517													1	9980	420
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ΕP	9753	48			B1		2004	1110									
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                         A2
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PRIORITY APPLN. INFO.:
                                            EP 1997-201180
                                                                A 19970422
                                                                W 19980420
                                            WO 1998-EP2361
                                            US 1999-403139
                                                                A1 19991014
                                            US 2002-117899
                                                                A1 20020408
                                            US 2002-303494
                                                                B1 20021125
ED
     Entered STN: 09 Nov 1998
     The invention pertains to a pharmaceutical dosage unit, such as a tablet
AB
     or a capsule, comprising an effective amount of tibolone (0.1-10 %
     by weight) and a starch-containing pharmaceutically acceptable carrier (also
     denoted as basic granules), wherein the carrier contains 10-100% by weight of
     the starch. Thus, a more stable tibolone formulation is
     obtained, allowing dry storage and lower doses of active ingredient.
     Thus, basic granules contained tibolone 0.3, HPC 1.95, corn
     starch 32.5, Mg stearate 0.325, and lactose to 65 mg.
ENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
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              2 S E2-E3
     FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005
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                E TIBOLONE/BI
            359 S E2-E4
L3
                E TIBOLONE/CT
L4
            430 S L2 OR L3
         147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L5
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L6
          78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L7
          61648 S LACTOSE OR DISACCHARIDE?
L8
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L9
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                E VD HURK T/AU
                E VAN DEN HURK T/AU
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L18 97 S E	E3-E6, E23 ROVERS A/AU		
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FILE 'BIOSIS' ENTERED AT 12:29:23 ON 25 MAR 2005 Copyright (c) 2005 The Thomson Corporation

FILE 'CAPLUS' ENTERED AT 12:29:23 ON 25 MAR 2005 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

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FILE 'EMBASE' ENTERED AT 12:29:23 ON 25 MAR 2005 COPYRIGHT (C) 2005 Elsevier Inc. All rights reserved.

FILE 'WPIDS' ENTERED AT 12:29:23 ON 25 MAR 2005 COPYRIGHT (C) 2005 THE THOMSON CORPORATION

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L26 2401 TIBOLONE? OR LIVIAL? OR LIVIELLA? OR "ORG OD14" OR "ORG-OD14" OR "ORG-OD14"

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        276310 STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
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     FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005
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L2
                E TIBOLONE/BI
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            359 S E2-E4
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L4
L5
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L6
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L32
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L32 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN
                         2004:1036906 CAPLUS
ACCESSION NUMBER:
                         142:11606
DOCUMENT NUMBER:
                         Immediate-release pharmaceutical dosage form
TITLE:
                         comprising a tibolone polymorph
                         De Haan, Pieter
INVENTOR(S):
                         Akzo Nobel N.V., Neth.
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 32 pp.
SOURCE:
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
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PATENT INFORMATION:
                                DATE
                                            APPLICATION NO.
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     WO 2004103378
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                                         WO 2004-EP50852
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PRIORITY APPLN. INFO.:
                                            EP 2003-101500
                                                              A 20030523
                                                               W 20040519
                                            WO 2004-EP50852
     Entered STN: 03 Dec 2004
ED
     An immediate-release pharmaceutical dosage form comprises a
AB
     tibolone polymorph and excipients, wherein the polymorph has a
     mean particle size as defined in the specification of below 22.8 \mu m in
     the dosage form. Thus, an immediate-release tablet formulation contained
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tibolone 2.5, potato starch 10, ascorbyl palmitate 0.2, Mg stearate 0.5, and lactose 86.8 mg.

REFERENCE COUNT:

5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L32 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

2004:453011 CAPLUS

DOCUMENT NUMBER:

140:429053

TITLE:

Pharmaceutical tablets containing tibolone

and a coating

INVENTOR(S):

De Haan, Pieter; Van Lare, Cornelis Elisabeth Johannus

PATENT ASSIGNEE(S):

Akzo Nobel N.V., Neth. PCT Int. Appl., 20 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	PATENT NO.				KIND DATE			APPLICATION NO.					DATE					
WO	2004	0455	87		A1	- :	2004	0603	1	WO 2	003-	EP50	829		2	0031	113	
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		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	KP,	KR,	ΚZ,	LC,	
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	ΝI,	NO,	
		NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,	TJ,	
		TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	zw		
	RW:	BW,	GH,	GM,	ΚE,	LS,	MW,	ΜZ,	SD,	SL,	SZ,	ΤZ,	UG,	ZM,	ZW,	AM,	ΑZ,	
		BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	
		ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	
		TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG
PRIORITY	PRIORITY APPLN. INFO.:								EP 2002-79920				0	i	A 20021120			
								US 2002-427740P P 2002			0021	120						

ED Entered STN: 04 Jun 2004

AΒ The present invention relates to a stabilized pharmaceutical tablet comprising an amount of from 0.1 to 10% by weight of tibolone provided with a coating, such as a film coating, a sugar coating, a sugar film coating or a wrap coating. The invention further relates to the use of a coating for stabilizing a pharmaceutical tablet comprising tibolone. Preferably, the coating is a sugar coating or a sugar film coating. Preferably, the sugar comprises sucrose. These tablets were shown to have a lower amount of OM38, i.e. an isomerization degradation product of tibolone, after storage as compared to uncoated tibolone tablets. For example, a tablet was formulated containing tibolone, basic granulates (starch/lactose), ascorbyl palmitate, and Mg stearate and a film-coated with a composition containing hydroxypropyl Me cellulose, polyethylene glycol, titania, and talc, further coated with a composition containing arabic gum, saccharose, Ca carbonate, talc, titania, polyethylene glycol, glucose syrup, glycerol, and china clay.

L32 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER:

1998:708947 CAPLUS

DOCUMENT NUMBER:

129:335779

TITLE:

SOURCE:

Stabilized tibolone compositions

INVENTOR(S):

Haan De, Pieter; Rovers, Adrianus Cornelis Petrus; Lambregts-De Hurk, Theodora Antonia Maria; Zwinkels,

Jocominus Antonius Maria; Morita, Ryoichi

PATENT ASSIGNEE(S):

Akzo Nobel N.V., Neth. PCT Int. Appl., 21 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PAT	rent :	NO.			KIN	D	DATE			API	PLI	CAT	ION	NO.			DATE	
	WO	9847				A1		1998	1029		WO	19	998-1	EP23	61			19980	1420
		W:	AM,	AU,	BB,	BG,	BR,	CA,	CN,	CZ,	E	Ξ,	GE,	HU,	ID,	IS,	KE	, KG,	KP,
			KR,	KZ,	LK,	LR,	LT,	LV,	MD,	MG,	M	N,	MX,	NO,	NZ,	PL,	RO	, SD,	SG,
																		, KG,	
					TJ,			·	•				,	•	•	•			•
		RW:	GH,	GM,	KE,	LS,	MW,	SD,	SZ,	UG,	ZV	N,	AT,	BE,	CH,	CY,	DE	, DK,	ES.
																		, CG,	
								NE,					•	•	•	•		••	•
	IL	1239				A1		2004					998-	1239	84			19980	407
	ZA	9803	169			Α		1998	1020									19980	415
	CA	2288	070			AA		1998	1029		CA	19	998-	2288	070			19980	420
	ΑU	9880	146			A1		1998						3014				19980	420
	ΑU	7420	37			R7		2001	1213										
	EΡ	9753	48			A1		2000	0202		ΕP	19	998-	9282	14			19980	420
	ΕP	9753	48			B1		2004	1110										
		R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GF	٦,	IT,	LI,	LU,	NL,	SE	, MC,	PT,
			ΙE,	FI															
		9902				Т2		2000						9902				19980	
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		2818				E		2004						9282				19980	
		1031				A2		1998						1123				19980	
		5707				В		2004							7211			19980	
		2001		13		A1		2001			US	19	99-4	1031	39			19991	014
		6399				B2		2002											
		9905				A		1999	-					5127				19991	
		9909				A		2000						9676				19991	
		6514		0.1		B1		2003						11789				20020	
		2003				A1		2003						3034				20021	
		2004				A1		2004						75468				20040	
		2004				A1		2004						75473				20040	
DDTA		2004				A1		2004) 122					7547				20040	
PRIO	KILI	APP:	LN.	INFO	. :							_		20118				19970	
														EP23				19980	
														10313				19991	
														1789				20020	
ED	Ent	bered	сти	. 0	a Not	, 100	38				US	20	102-3	30349	14		ВI	20021	125

ED Entered STN: 09 Nov 1998

AB The invention pertains to a pharmaceutical dosage unit, such as a tablet or a capsule, comprising an effective amount of tibolone (0.1-10 % by weight) and a starch-containing pharmaceutically acceptable carrier (also denoted as basic granules), wherein the carrier contains 10-100% by weight of the starch. Thus, a more stable tibolone formulation is obtained, allowing dry storage and lower doses of active ingredient. Thus, basic granules contained tibolone 0.3, HPC 1.95, corn starch 32.5, Mg stearate 0.325, and lactose to 65 mg.

REFERENCE COUNT:

5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> dup rem 133
PROCESSING COMPLETED FOR L33
L34 5 DUP REM L33 (1 DUPLICATE REMOVED)

ANSWER '1' FROM FILE BIOSIS ANSWERS '2-4' FROM FILE CAPLUS ANSWER '5' FROM FILE WPIDS

=> s 134 not 132

L35 2 L34 NOT L32

=> d 135 1-2 ibib ed abs

L35 ANSWER 1 OF 2 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation on STN

ACCESSION NUMBER: 2002:378050 BIOSIS DOCUMENT NUMBER: PREV200200378050

TITLE: Stabilized tibolone compositions.

AUTHOR(S): de Haan, Pieter [Inventor, Reprint author]; Lambregts v.d. Hurk, Theodora Antonia Maria [Inventor]; Morita, Ryoichi

[Inventor]; Rovers, Adrianus Cornelis Petrus [Inventor];

Zwinkels, Jocominus Antonius Maria [Inventor]

CORPORATE SOURCE: Oss, Netherlands

ASSIGNEE: Akzo Nobel NV, Arnhem, Netherlands

PATENT INFORMATION: US 6399594 June 04, 2002

SOURCE: Official Gazette of the United States Patent and Trademark

Office Patents, (June 4, 2002) Vol. 1259, No. 1. http://www.uspto.gov/web/menu/patdata.html. e-file.

CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE: Patent LANGUAGE: English

ENTRY DATE: Entered STN: 10 Jul 2002

Last Updated on STN: 10 Jul 2002

ED Entered STN: 10 Jul 2002

Last Updated on STN: 10 Jul 2002

AB The invention pertains to a pharmaceutical dosage unit, such as a tablet or a capsule, comprising an effective amount of tibolone (generally of from 0.1 to 10% by weight) and a starch-containing pharmaceutically acceptable carrier (also denoted as basic granulate), wherein the carrier contains of from 10 to 100% by weight of the starch. Thus a more stable tibolone formulation is

obtained, allowing dry storage and lower doses of active ingredient.

L35 ANSWER 2 OF 2 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2003-393472 [37] WPIDS

DOC. NO. CPI: C2003-104580

TITLE: Tibolone formulations with improved stability,

useful as a gonadomimetic and immunomodulator, and for

inhibiting ovulation.

DERWENT CLASS: A96 B01

INVENTOR(S): BRENNAN, T; WOOLFE, A J

PATENT ASSIGNEE(S): (BREN-I) BRENNAN T; (WOOL-I) WOOLFE A J; (IVAX-N) IVAX

CORP; (NORT-N) NORTON HEALTHCARE LTD

COUNTRY COUNT: 101

PATENT INFORMATION:

PATENT NO	KIND DATE	WEEK	LA	PG

WO 2003032924 A2 20030424 (200337) * EN 19

RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR IE IT KE LS LU

MC MW MZ NL OA PT SD SE SK SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK

DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR

KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI SK SL TJ TM TN TR TT TZ UA UG UZ VN YU ZA ZM ZW

US 2003134832 A1 20030717 (200348)

EP 1444247 A2 20040811 (200452) EN

R: AL AT BE BG CH CY CZ DE DK EE ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI SK TR

AU 2002347973 A1 20030428 (200460)

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2003032924	A2	WO 2002-US33526	20021018
US 2003134832	Al Provisional	US 2001-350121P US 2002-273455	20011102 20021018
EP 1444247	A2	EP 2002-784184 WO 2002-US33526	20021018 20021018
AU 2002347973	A1	AU 2002-347973	20021018

FILING DETAILS:

PATENT NO	KIND	PATENT NO
EP 1444247	A2 Based on	WO 2003032924
AU 2002347973	A1 Based on	WO 2003032924

PRIORITY APPLN. INFO: US 2001-350121P 20011102; GB 20011018 2001-25061

ED 20030612

AN 2003-393472 [37] WPIDS

AB WO2003032924 A UPAB: 20030612

> NOVELTY - A composition comprises tibolone, one or more excipients, and a pH adjusting agent.

ACTIVITY - Immunomodulator; Gynecological.

MECHANISM OF ACTION - Gonadomimetic.

USE - Tibolone has gonadomimetic, ovulation-inhibiting and immuno-modulatory action.

ADVANTAGE - The inclusion of a pH adjusting agent improves the stability of the compositions, even those with a low tibolone content, and retains their solubility. Dwq.0/2

=> d his

(FILE 'HOME' ENTERED AT 12:12:09 ON 25 MAR 2005)

FILE 'REGISTRY' ENTERED AT 12:12:34 ON 25 MAR 2005 E TIBOLONE/CN

2 S E2-E3 L1

FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005

L2 386 S L1 E TIBOLONE/BI 359 S E2-E4 L3E TIBOLONE/CT 430 S L2 OR L3 L4

147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE L5 L6

348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP

78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC L7

61648 S LACTOSE OR DISACCHARIDE? r_8

5 S L4 AND L5 AND (L6 OR L7) L9

L10 3 S TIBOLONE? AND STARCH? 0 S L10 NOT L9 L11

7 S L4 AND L8 L12 2 S L12 NOT L9 L13

E HAAN P/AU L14 3 S E3, E5, E7

E DE HAAN P/AU L15 96 S E3-E9, E17-E18

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E LAMBREGTS T/AU
L16
              1 S E4
                E HURK T/AU
                E VD HURK T/AU
                E VAN DEN HURK T/AU
L17
              1 S E4
                E MORITA R/AU
             97 S E3-E6, E23
1.18
                E ROVERS A/AU
              1 S E4
1.19
                E ZWINKELS J/AU
              3 S E3, E8
L20
             99 S L14 OR L15
L21
L22
            198 S L21 OR L16 OR L17 OR L18 OR L19 OR L20
L23
              4 S L22 AND TIBOLONE?
     FILE 'MEDLINE, BIOSIS, CAPLUS, EMBASE, WPIDS' ENTERED AT 12:29:23 ON 25
     MAR 2005
L24
           8818 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR BOLTIN? OR "ORG OD14"
           2399 S L24 NOT BOLTIN?
L25
           2401 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR "ORG OD14" OR "ORG OD14"
L26
         276310 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L27
L28
         693483 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
         176163 S (ASCORBYL PALMITATE?) OR "6-0-PALMITOYLASCORBIC ACID" OR "ASC
L29
         139122 S LACTOSE OR DISACCHARIDE?
L30
              3 S L26 AND L27 AND (L28 OR L29)
L31
              3 DUP REM L31 (0 DUPLICATES REMOVED)
L32
              6 S TIBOLONE? AND STARCH?
L33
L34
              5 DUP REM L33 (1 DUPLICATE REMOVED)
L35
              2 S L34 NOT L32
=> s 126 and 130
             8 L26 AND L30
L36
=> dup rem 136 ·
PROCESSING COMPLETED FOR L36
1.37
              5 DUP REM L36 (3 DUPLICATES REMOVED)
                ANSWER '1' FROM FILE MEDLINE
                ANSWERS '2-5' FROM FILE CAPLUS
=> s 137 not 134
             2 L37 NOT L34
L38
=> d 128 1-2 ibib ed abs
L28 ANSWER 1 OF 693483
                            MEDLINE on STN
ACCESSION NUMBER:
                    2005153240
                                    IN-PROCESS
                    PubMed ID: 15785787
DOCUMENT NUMBER:
TITLE:
                    [Effects of psychiatric drugs on the fetus and newborn.
                    Consequences of the treatment of psychiatric disorders
                    during pregnancy and lactation].
                    Efectos de los psicofarmacos en el feto y el recien nacido.
                    Consecuencias del tratamiento de los trastornos
                    psiquiatricos durante el embarazo y la lactancia.
AUTHOR:
                    Mazaira Silvina
CORPORATE SOURCE:
                    Facultad de Medicina, Universidad de Buenos Aires,
                    Argentina.. smazaira@arnet.com.ar
SOURCE:
                    Vertex (Buenos Aires, Argentina), (2005 Jan) 16 (59) 35-42.
                    Journal code: 9440528. ISSN: 0327-6139.
PUB. COUNTRY:
                    Argentina
DOCUMENT TYPE:
                    Journal; Article; (JOURNAL ARTICLE)
LANGUAGE:
FILE SEGMENT:
                    NONMEDLINE; IN-DATA-REVIEW; IN-PROCESS; NONINDEXED;
                    Priority Journals
```

ENTRY DATE: Entered STN: 20050324

Last Updated on STN: 20050324

ED Entered STN: 20050324

Last Updated on STN: 20050324

AB The management of mental disorders during pregnancy and the postpartum period confronts the psychiatrist to a complex clinical situation. The cero risk utopia should be changed for a more realistic approach which considers the risks of having a conservative conduct (assuming the consequences of the untreated mental disorders) or making a psychopharmacological intervention knowing the expectable effects on the fetus and the offspring. The antidepressants and the conventional antipsychotics seem to be safe drugs in the first trimester of pregnancy. The benzodiazepines and the mood stabilizers are linked with the appearance of teratogenic malformations after the first trimester exposure. In the present article, the pharmacokinetic peculiarities of the pregnancy and the newborn and the general considerations of the psychopharmacological management of this kind of patients are described. At last, there is a review of the literature about the effects of the different groups of psychiatric drugs on the fetal growth, their passage to maternal milk and their impact on the newborn.

L28 ANSWER 2 OF 693483 MEDLINE on STN ACCESSION NUMBER: 2005153188 IN-PROCESS

DOCUMENT NUMBER: PubMed ID: 15647597

TITLE: Activation of Mitogen Activated Protein Kinases by PCB126

(3,3',4,4',5-Pentachlorobiphenyl) in HepG2 Cells.

AUTHOR: Song Min Ok; Freedman Jonathan H

CORPORATE SOURCE: Nicholas School of the Environment and Earth Sciences, Duke

University, Durham, North Carolina 27708-0328.

SOURCE: Toxicological sciences: an official journal of the Society

of Toxicology, (2005 Apr) 84 (2) 308-18. Electronic

Publication: 2005-01-12.

Journal code: 9805461. ISSN: 1096-6080.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: NONMEDLINE; IN-DATA-REVIEW; IN-PROCESS; NONINDEXED;

Priority Journals

ENTRY DATE: Entered STN: 20050324

Last Updated on STN: 20050324

ED Entered STN: 20050324

Last Updated on STN: 20050324

Polychlorinated biphenyls (PCBs) are persistent organic pollutants. AΒ Exposure to PCB126 (3,3',4,4',5-pentachlorobiphenyl), a non-ortho-chlorinated, coplanar congener has been correlated with the production of reactive oxygen species in vivo. In this report, we show that treatment of HepG2 cells with PCB126 significantly increases oxidative-stress-responsive transcription. In addition, PCB126-induced transcription is enhanced in cells depleted of glutathione. Exposure to PCB126 induces a cellular stress response in HepG2 cells that results in a significant increase in the activity of the mitogen activated protein kinases: extracellular signal regulated kinases 1/2 and p38. PCB126 exposure also causes an increase in c-Jun phosphorylation. These results suggest a model for PCB126 toxicity in which PCB126 exposure induces oxidative stress that causes an increase in mitogen-activated protein kinase (MAPK) activities and a subsequent increase in c-Jun phosphorylation. Activation of c-Jun results in enhanced activating protein-1 (AP-1) activity, which leads to elevated expression of antioxidant responsive element (ARE)/AP-1-dependent genes.

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FILE 'REGISTRY' ENTERED AT 12:12:34 ON 25 MAR 2005
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L1
              2 S E2-E3
     FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005
            386 S L1
L2
                E TIBOLONE/BI
L3
            359 S E2-E4
                E TIBOLONE/CT
L4
            430 S L2 OR L3
         147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L5
         348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
L6
          78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L7
          61648 S LACTOSE OR DISACCHARIDE?
rac{1}{8}
              5 S L4 AND L5 AND (L6 OR L7)
L9
              3 S TIBOLONE? AND STARCH?
L10
              0 S L10 NOT L9
L11
              7 S L4 AND L8
L12
L13
              2 S L12 NOT L9
                E HAAN P/AU
L14
              3 S E3, E5, E7
                E DE HAAN P/AU
L15
             96 S E3-E9, E17-E18
                E LAMBREGTS T/AU
L16
              1 S E4
                E HURK T/AU
                E VD HURK T/AU
                E VAN DEN HURK T/AU
L17
              1 S E4
                E MORITA R/AU
             97 S E3-E6, E23
L18
                E ROVERS A/AU
L19
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                E ZWINKELS J/AU
L20
              3 S E3, E8
L21
             99 S L14 OR L15
L22
            198 S L21 OR L16 OR L17 OR L18 OR L19 OR L20
L23
              4 S L22 AND TIBOLONE?
     FILE 'MEDLINE, BIOSIS, CAPLUS, EMBASE, WPIDS' ENTERED AT 12:29:23 ON 25
     MAR 2005
L24
           8818 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR BOLTIN? OR "ORG OD14"
L25
           2399 S L24 NOT BOLTIN?
           2401 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR "ORG OD14" OR "ORG OD14"
L26
         276310 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L27
L28
         693483 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
L29
         176163 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
         139122 S LACTOSE OR DISACCHARIDE?
L30
L31
              3 S L26 AND L27 AND (L28 OR L29)
L32
              3 DUP REM L31 (0 DUPLICATES REMOVED)
L33
              6 S TIBOLONE? AND STARCH?
L34
              5 DUP REM L33 (1 DUPLICATE REMOVED)
L35
              2 S L34 NOT L32
L36
              8 S L26 AND L30
L37
              5 DUP REM L36 (3 DUPLICATES REMOVED)
L38
              2 S L37 NOT L34
=> s "boltin"
L39
             5 "BOLTIN"
=> s 139 and starch?
L40
             0 L39 AND STARCH?
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L1
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     FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005
L2
            386 S L1
                E TIBOLONE/BI
            359 S E2-E4
L3
                E TIBOLONE/CT
            430 S L2 OR L3
L4
L5
         147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
         348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
L6
L7
          78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
          61648 S LACTOSE OR DISACCHARIDE?
L8
L9
              5 S L4 AND L5 AND (L6 OR L7)
L10
              3 S TIBOLONE? AND STARCH?
              0 S L10 NOT L9
L11
L12
              7 S L4 AND L8
L13
              2 S L12 NOT L9
                E HAAN P/AU
L14
              3 S E3, E5, E7
                E DE HAAN P/AU
L15
             96 S E3-E9, E17-E18
                E LAMBREGTS T/AU
              1 S E4
L16
                E HURK T/AU
                E VD HURK T/AU
                E VAN DEN HURK T/AU
L17
              1 S E4
                E MORITA R/AU
             97 S E3-E6, E23
L18
                E ROVERS A/AU
L19
              1 S E4
                E ZWINKELS J/AU
              3 S E3, E8
L20
L21
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L22
L23
              4 S L22 AND TIBOLONE?
     FILE 'MEDLINE, BIOSIS, CAPLUS, EMBASE, WPIDS' ENTERED AT 12:29:23 ON 25
     MAR 2005
L24
           8818 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR BOLTIN? OR "ORG OD14"
L25
           2399 S L24 NOT BOLTIN?
           2401 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR "ORG OD14" OR "ORG OD14"
L26
         276310 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L27
         693483 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
L28
         176163 S (ASCORBYL PALMITATE?) OR "6-0-PALMITOYLASCORBIC ACID" OR "ASC
L29
         139122 S LACTOSE OR DISACCHARIDE?
L30
L31
              3 S L26 AND L27 AND (L28 OR L29)
              3 DUP REM L31 (0 DUPLICATES REMOVED)
L32
              6 S TIBOLONE? AND STARCH?
L33
              5 DUP REM L33 (1 DUPLICATE REMOVED)
L34
              2 S L34 NOT L32
L35
L36
              8 S L26 AND L30
              5 DUP REM L36 (3 DUPLICATES REMOVED)
L37
              2 S L37 NOT L34
L38
             5 S "BOLTIN"
L39
L40
              0 S L39 AND STARCH?
```

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
CONNECT CHARGES	27.27	38.71
NETWORK CHARGES SEARCH CHARGES	0.90 139.86	2.70 300.26
DISPLAY CHARGES	14.19	43.34
FULL ESTIMATED COST	182.22	385.01
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	-2.19	-10.22

IN FILE 'MEDLINE, BIOSIS, CAPLUS, EMBASE, WPIDS' AT 12:38:20 ON 25 MAR 2005

=> file stngudie 'STNGUDIE' IS NO

'STNGUDIE' IS NOT A VALID FILE NAME
SESSION CONTINUES IN FILE 'MEDLINE, BIOSIS, CAPLUS, EMBASE, WPIDS'
Enter "HELP FILE NAMES" at an arrow prompt (=>) for a list of files
that are available. If you have requested multiple files, you can
specify a corrected file name or you can enter "IGNORE" to continue
accessing the remaining file names entered.

=> file stnguide		
COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	186.22	389.01
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
	ENTRY	SESSION
CA SUBSCRIBER PRICE	-2.19	-10.22

FILE 'STNGUIDE' ENTERED AT 12:38:54 ON 25 MAR 2005
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COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY, JAPAN SCIENCE
AND TECHNOLOGY CORPORATION, AND FACHINFORMATIONSZENTRUM KARLSRUHE

FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: Mar 18, 2005 (20050318/UP).

=> file registry		
COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	0.18	389.19
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
	ENTRY	SESSION
CA SUBSCRIBER PRICE	0.00	-10.22

FILE 'REGISTRY' ENTERED AT 12:40:55 ON 25 MAR 2005 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2005 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 24 MAR 2005 HIGHEST RN 847222-24-6 DICTIONARY FILE UPDATES: 24 MAR 2005 HIGHEST RN 847222-24-6

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 18, 2005

Please note that search-term pricing does apply when

conducting SmartSELECT searches.

E2

E3

E4

***************** The CA roles and document type information have been removed from * the IDE default display format and the ED field has been added, effective March 20, 2005. A new display format, IDERL, is now available and contains the CA role and document type information. ***************** Crossover limits have been increased. See HELP CROSSOVER for details. Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html => e potato starch/cn POTATO SPINDLE TUBER VIROID RNA-BINDING PROTEIN VIRP1 (TOMAT E1O CLONE VIRP1C)/CN E2 POTATO SPINDLE TUBER VIROID RNA-BINDING PROTEIN VIRP1 (TOMAT O CLONE VIRP1D)/CN F.3 1 --> POTATO STARCH/CN E4 1 POTATO STARCH BENZOATE ISOSTEARATE/CN POTATO, EXT./CN E5 1 E6 1 POTATO, EXT., SAPOND./CN POTATO, SOLANUM TUBEROSUM AEGROTANS, EXT./CN E7 1 E8 1 POTAVESCENT/CN E9 POTB (MYCOPLASMA GALLISEPTICUM STRAIN RLOW GENE POTB)/CN 1 1 POTB (PASTEURELLA MULTOCIDA STRAIN IL1403 CLONE PM70 GENE PO E10 TB)/CN 1 POTB (STREPTOCOCCUS PYOGENES STRAIN MGAS10394)/CN E11 E12 POTB PROTEIN (MANNHEIMIA SUCCINICIPRODUCENS STRAIN MBEL55E G ENE POTB)/CN => s e3 L41 1 "POTATO STARCH"/CN => e corn starch/cn CORN P 04/CN E1 1 E2 CORN PROTEIN HYDROLYZATES/CN E3 1 --> CORN STARCH/CN E4CORN STARCH ACETATE BENZOATE/CN E5 CORN STARCH-FORMALDEHYDE-HEXAMETHYLENETETRAMINE-PHENOL COPOL YMER/CN E6 1 CORN STARCH-HYDRAN HW 112-SUMIKAFLEX 410-YODO SOL YL 2 COPOL YMER/CN E7 CORN STEEP WATER/CN E8 CORN STRAIN B73 CHROMOSOME 9S BRONZE GENE-CONTAINING REGION/ 1 CORN SUGAR/CN E9 1 CORN SYRUP/CN E10 E11 1 CORN Y/CN CORN, BRAN/CN E12 => s e3 L42 1 "CORN STARCH"/CN => e wheat starch/cn E1 WHEAT GERM-OIL GLYCERIDES/CN 1

WHEAT GLUTEN HYDROLYZATES/CN

0 --> WHEAT STARCH/CN

WHEAT VIM/CN

E5 E6 E7 E8 E9 E10 E11	1 1 1 1 1 1 1	WHEAT, EXT./CN WHEAT, EXT., ET ESTER/CN WHEAT, EXT., HYDROLYZED/CN WHEAT, FLOUR/CN WHEAT-OIL FATTY ACIDS/CN WHEATLEYITE/CN WHEATLEYITE (NA2(CU(C2O4)2).2H2O)/CN WHEATWIN1 (TRITICUM AESTIVUM VARIETY SAN PASTORE ASPARAGINE-CONTAINING ISOFORM N-TERMINAL BLOCKAGE)/CN
=> e modifie		
E1 E2	5 1	MODIFIED PEPTIDE (SYNTHETIC EBV FUSION INHIBITOR)/CN MODIFIED PROSECRETASE B (SYNTHETIC PLASMID PET11A-BACE-
E3	0 .	PRO33LYS FRAGMENT)/CN
E4		MODIFIED STARCH/CN
	1	MODIFIED VASCO X2/CN
E5	1	MODIFIER A/CN
E6	1	MODIFIER KS 4930/CN
E7	1	MODIFIER OF CITRATE LYASE PROTEIN (SALMONELLA ENTERICA TYPHI MURIUM STRAIN LT2; SGSC 1412; ATCC 700720 GENE CITG)/CN
E8	1	MODIFIER OF CITRATE LYASE PROTEIN (SALMONELLA ENTERICA TYPHI
		MURIUM STRAIN LT2; SGSC 1412; ATCC 700720 GENE CITG2)/CN
E9	1	MODIFIER PROTEIN (YERSINIA PESTIS STRAIN CO92 GENE YPO3046)/CN
E10	1	MODIFIER PROTEIN OF MAJOR AUTOLYSIN LYTC (CWBP76) (BACILLUS
		LICHENIFORMIS STRAIN ATCC 14580 GENE LYTB)/CN
E11	1	MODIFIER PROTEIN OF MAJOR AUTOLYSIN LYTC (CWBP76) (BACILLUS
		SUBTILIS GENE LYTB)/CN
E12	1	MODIFILAN/CN
=> e agglome		
E1	1	AGGLOFROID 009/CN
E2	1	AGGLOFROID 313E/CN
E3		AGGLOMERATED STARCH/CN
E4	1	AGGLOMERIN A/CN
E5	1	AGGLOMERIN B/CN
E6	1	AGGLOMERIN C/CN
E7	1	AGGLOMERIN D/CN
E8	3	AGGLOMERONE/CN
E9	1	AGGLOMERONE IMINE/CN
E10	1	AGGLOMERONE, COPPER DERIV./CN
E11	1	AGGLUCETIN/CN
E12	1	AGGLUCETIN (AGKISTRODON ACUTUS SUBUNIT A1 PRECURSOR)/C
		N
=> e granula		
E1	1	GRANULAR BLUE/CN
E2	1	GRANULAR EXPLOSIVES/CN
E3		GRANULATED STARCH/CN
E4	1	GRANULATED SUGAR/CN
E5	1	GRANULATIMI DE/CN
E6	1	GRANULATIN/CN
E7	1	GRANULATOSIDE A/CN
E8	1	GRANULATOSIDE B/CN
E9	1	GRANULE ANTIGEN PROTEIN GRA6 (TOXOPLASMA GONDII STRAIN BEVER LEY GENE GRA6)/CN
E10	1	GRANULE ANTIGEN PROTEIN GRA6 (TOXOPLASMA GONDII STRAIN CASTE LLS GENE GRA6)/CN
E11	1	GRANULE ANTIGEN PROTEIN GRA6 (TOXOPLASMA GONDII STRAIN FOU G
E12	1	ENE GRA6)/CN
E12	1	GRANULE ANTIGEN PROTEIN GRA6 (TOXOPLASMA GONDII STRAIN M7741 GENE GRA6)/CN
		J. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1.

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	10.06	399.25
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	0.00	-10.22

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FILE COVERS 1907 - 25 Mar 2005 VOL 142 ISS 14 FILE LAST UPDATED: 24 Mar 2005 (20050324/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d his

L17

(FILE 'HOME' ENTERED AT 12:12:09 ON 25 MAR 2005)

E VAN DEN HURK T/AU

1 S E4

FILE 'REGISTRY' ENTERED AT 12:12:34 ON 25 MAR 2005 E TIBOLONE/CN

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L1
              2 S E2-E3
     FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005
L2
            386 S L1
                E TIBOLONE/BI
            359 S E2-E4
L3
                E TIBOLONE/CT
            430 S L2 OR L3
L4
L5
         147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
         348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
L6
          78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L7
          61648 S LACTOSE OR DISACCHARIDE?
r_8
L9
              5 S L4 AND L5 AND (L6 OR L7)
              3 S TIBOLONE? AND STARCH?
L10
              0 S L10 NOT L9
L11
              7 S L4 AND L8
L12
L13
              2 S L12 NOT L9
                E HAAN P/AU
L14
              3 S E3, E5, E7
                E DE HAAN P/AU
L15
             96 S E3-E9, E17-E18
                E LAMBREGTS T/AU
              1 S E4
L16
                E HURK T/AU
                E VD HURK T/AU
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97 S E3-E6, E23
L18
                E ROVERS A/AU
L19
              1 S E4
                E ZWINKELS J/AU
L20
              3 S E3, E8
L21
             99 S L14 OR L15
L22
            198 S L21 OR L16 OR L17 OR L18 OR L19 OR L20
L23
              4 S L22 AND TIBOLONE?
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     MAR 2005
           8818 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR BOLTIN? OR "ORG OD14"
L24
           2399 S L24 NOT BOLTIN?
L25
           2401 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR "ORG OD14" OR "ORG OD14"
L26
         276310 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L27
         693483 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
L28
         176163 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L29
         139122 S LACTOSE OR DISACCHARIDE?
L30
              3 S L26 AND L27 AND (L28 OR L29)
L31
L32
              3 DUP REM L31 (0 DUPLICATES REMOVED)
              6 S TIBOLONE? AND STARCH?
L33
              5 DUP REM L33 (1 DUPLICATE REMOVED)
L34
              2 S L34 NOT L32
L35
              8 S L26 AND L30
L36
L37
              5 DUP REM L36 (3 DUPLICATES REMOVED)
              2 S L37 NOT L34
L38
              5 S "BOLTIN"
L39
              0 S L39 AND STARCH?
L40
     FILE 'STNGUIDE' ENTERED AT 12:38:54 ON 25 MAR 2005
     FILE 'REGISTRY' ENTERED AT 12:40:55 ON 25 MAR 2005
                E POTATO STARCH/CN
L41
              1 S E3
                E CORN STARCH/CN
L42
              1 S E3
                E WHEAT STARCH/CN
                E MODIFIED STARCH/CN
                E AGGLOMERATED STARCH/CN
                E GRANULATED STARCH/CN
     FILE 'CAPLUS' ENTERED AT 12:42:18 ON 25 MAR 2005
=> s 14 and (15 or 141 or 142)
         72830 L41
         72830 L42
L43
             6 L4 AND (L5 OR L41 OR L42)
=> d 143 1-6 ibib ed abs
L43 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
                         2004:1036906 CAPLUS
DOCUMENT NUMBER:
                          142:11606
TITLE:
                         Immediate-release pharmaceutical dosage form
                         comprising a tibolone polymorph
INVENTOR(S):
                         De Haan, Pieter
PATENT ASSIGNEE(S):
                         Akzo Nobel N.V., Neth.
                         PCT Int. Appl., 32 pp.
SOURCE:
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
                         English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
                         1
PATENT INFORMATION:
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E MORITA R/AU

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PATENT NO.
                                  KIND
                                             DATE
                                                           APPLICATION NO.
                                                                                              DATE
                                                             -----
                                                                                              _____
       WO 2004103378
                                   A1
                                             20041202
                                                          WO 2004-EP50852
                                                                                              20040519
            W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
                  CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
                  LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
                  NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
                  TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
            RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
                  SN, TD, TG
       CA 2473718
                                             20041123
                                                             CA 2004-2473718
                                    AΑ
                                                                                              20040519
       EP 1499278
                                                             EP 2004-737220
                                             20050126
                                    A1
                                                                                              20040519
            R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR
PRIORITY APPLN. INFO.:
                                                              EP. 2003-101500
                                                                                       A 20030523
                                                                                         W 20040519
                                                              WO 2004-EP50852
ED
       Entered STN: 03 Dec 2004
AB
      An immediate-release pharmaceutical dosage form comprises a
       tibolone polymorph and excipients, wherein the polymorph has a
      mean particle size as defined in the specification of below 22.8 μm in
       the dosage form. Thus, an immediate-release tablet formulation contained
       tibolone 2.5, potato starch 10, ascorbyl
       palmitate 0.2, Mg stearate 0.5, and lactose 86.8 mg.
REFERENCE COUNT:
                                           THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
                                   5
                                           RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L43 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2005 ACS on STN
                                   2004:453011 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                                   140:429053
TITLE:
                                   Pharmaceutical tablets containing tibolone
                                   and a coating
INVENTOR(S):
                                   De Haan, Pieter; Van Lare, Cornelis Elisabeth Johannus
                                  Akzo Nobel N.V., Neth.
PATENT ASSIGNEE(S):
                                   PCT Int. Appl., 20 pp.
SOURCE:
                                   CODEN: PIXXD2
DOCUMENT TYPE:
                                   Patent
LANGUAGE:
                                   English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
      PATENT NO.
                                  KIND
                                             DATE
                                                             APPLICATION NO.
                                   ----
                                            -----
                                                             ------
                                                                                              _____
      WO 2004045587
                                   A1
                                                          WO 2003-EP50829
                                            20040603
                                                                                              20031113
           W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BE, BJ, CF, CG, CI, CM, GA, GN, GO, GW, MI, MR, NE, SN, TD,
                  TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD,
                                                             EP 2002-79920
                                                                                     A 20021120
P 20021120
PRIORITY APPLN. INFO.:
                                                             US 2002-427740P
```

ED Entered STN: 04 Jun 2004

AB The present invention relates to a stabilized pharmaceutical tablet comprising an amount of from 0.1 to 10% by weight of **tibolone** provided with a coating, such as a film coating, a sugar coating, a sugar

film coating or a wrap coating. The invention further relates to the use of a coating for stabilizing a pharmaceutical tablet comprising tibolone. Preferably, the coating is a sugar coating or a sugar film coating. Preferably, the sugar comprises sucrose. These tablets were shown to have a lower amount of OM38, i.e. an isomerization degradation product of tibolone, after storage as compared to uncoated tibolone tablets. For example, a tablet was formulated containing tibolone, basic granulates (starch/lactose), ascorbyl palmitate, and Mg stearate and a film-coated with a composition containing hydroxypropyl Me cellulose, polyethylene glycol, titania, and talc, further coated with a composition containing arabic qum, saccharose, Ca

talc, titania, polyethylene glycol, glucose syrup, glycerol, and china

L43 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1998:708947 CAPLUS

DOCUMENT NUMBER:

129:335779

TITLE:

Stabilized tibolone compositions

INVENTOR(S):

Haan De, Pieter; Rovers, Adrianus Cornelis Petrus; Lambregts-De Hurk, Theodora Antonia Maria; Zwinkels,

Jocominus Antonius Maria; Morita, Ryoichi

PATENT ASSIGNEE(S):

Akzo Nobel N.V., Neth. PCT Int. Appl., 21 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

WO 9847517	PAT	ENT 1	NO.			KIN	D	DATE		i	APPL	ICAT	ION I	NO.		D	ATE	
W: AM, AU, BB, BG, BR, CA, CN, CZ, EE, GE, HU, ID, IS, KE, KG, KP, KR, KZ, LK, LR, LT, LV, MD, MG, MN, MX, NO, NZ, PL, RO, SD, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ, VN, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG IL 123984 A1 20040927 IL 1998-123984 19980407 ZA 9803169 A1 19981020 ZA 1998-3169 19980415 CA 2288070 AA 19981020 ZA 1998-3169 19980415 CA 2288070 AA 19981113 AU 1998-80146 19980420 AU 742037 B2 20011213 EP 975348 B1 20041110 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI TR 9902530 T2 20000522 TR 1999-9902530 19980420 BR 9809268 A 20000627 BR 1998-9268 19980420 RU 2207133 C2 20030627 RU 1999-121407 19980420 RU 2207133 C2 20030627 RU 1998-112312 19980420 AT 281836 E 20041115 AT 1998-9268 19980420 AT 281836 E 20041115 AT 1998-9268 19980420 AT 281836 E 20041115 AT 1998-921407 19980420 AT 281836 E 20041115 AT 1998-921407 19980420 AT 281836 E 2004111 TW 1998-87107211 19980511 US 2001005513 A1 20010628 US 1999-403139 19991014 US 6399594 B2 20020604 NO 9905127 A 19991021 NO 1999-5127 19991021 US 6514958 B1 20030204 US 2002-117899 20020408	MO	9817	 517			Δ1		1998	1029	,	ฬก 1	998-1	EP23	61		1	9980	420
KR, KZ, LK, LR, LT, LV, MD, MG, MN, MX, NO, NZ, PL, RO, SD, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ, VN, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG IL 123984	""																	
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MD, RU, TJ, TM																		
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG IL 123984				•	•	•	,	,	,	,	,	,	•	,	,	,	,	,
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CA 2288070	ZA	9803	169			Α		1998	1020		ZA 1	998-	3169			1	9980	415
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US 2004142028 A1 20040722 US 2004-754732 20040108	US	2004	1420	28		A1		2004	0722	1	US 2	004-	7547	32		2	0040	108

US 2004142031 A1 20040722 US 2004-754733 20040108
PRIORITY APPLN. INFO.: EP 1997-201180 A 19970422
WO 1998-EP2361 W 19980420
US 1999-403139 A1 19991014
US 2002-117899 A1 20020408
US 2002-303494 B1 20021125

ED Entered STN: 09 Nov 1998

AB The invention pertains to a pharmaceutical dosage unit, such as a tablet or a capsule, comprising an effective amount of tibolone (0.1-10 % by weight) and a starch-containing pharmaceutically acceptable carrier (also denoted as basic granules), wherein the carrier contains 10-100% by weight of the starch. Thus, a more stable tibolone formulation is obtained, allowing dry storage and lower doses of active ingredient. Thus, basic granules contained tibolone 0.3, HPC 1.95, corn starch 32.5, Mg stearate 0.325, and lactose to 65 mg.

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L43 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1996:313906 CAPLUS

DOCUMENT NUMBER: 124:352708

TITLE: Solid pharmaceutical composition comprising an

excipient capable of binding water

INVENTOR(S): De Haan, Pieter; Poels-Janssen, Henrika Gerardina

Maria

PATENT ASSIGNEE(S): Akzo Nobel N.V., Neth.

SOURCE: Eur. Pat. Appl., 9 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PAT	TENT NO.		KIND	DATE	APPLICATION NO.	DATE
	707848 707848		A1 B1	19960424 20020109	EP 1995-202770	19951013
	R: AT,	BE, CH,	DE, D	K, ES, FR,	GB, GR, IE, IT, LI, NL,	PT, SE
CA	2159419		AA	19960418	CA 1995-2159419	19950928
IL	115445		A1	19990817	IL 1995-115445	19950928
BR	9504400		Α	19970527	BR 1995-4400	19951013
AT	211647		E	20020115	AT 1995-202770	19951013
PT	707848		${f T}$	20020628	PT 1995-202770	19951013
ES	2171498		Т3	20020916	ES 1995-202770	19951013
FI	9504905		Α	19960418	FI 1995-4905	19951016
AU	9534267		A1	19960502	AU 1995-34267	19951016
UA	688581		B2	19980312		
CN	1130064		Α	19960904	CN 1995-116176	19951016
CN	1112185		В	20030625		
HU	75247		A2	19970528	ни 1995-2985	19951016
HU	221427		В	20021028		
RU	2152802		C2	20000720	RU 1995-118099	19951016
	08268914		A2	19961015	JP 1995-268772	19951017
US	6187339		B1	20010213	US 1995-544042	19951017
	1001886		A1	20020531	нк 1998-101017	19980211
PRIORITY	APPLN.	INFO.:			EP 1994-203017	A 19941017

ED Entered STN: 30 May 1996

The invention concerns a solid pharmaceutical composition comprising less than 7% by weight of an oil or oily substance, a low dosage active ingredient, and a water-insol. noncrosslinked polymeric excipient capable of binding water and having a mean particle size greater than 150 μm . The invention offers a solution for obtaining drug-loaded carrier particles without the need of a granulation step by using the solid composition. The composition can

obtained by a simple procedure comprising mixing the water-insol. noncrosslinked polymeric excipient capable of binding water, preferably cellulose or starch, and the active ingredient, which is dissolved or dispersed in an oil or an oily substance, in an aqueous dispersion thereof, or in water. A tablet contained desogestrel 150, ethinyl estradiol 30 µg, Miglyol 812 1.3, water 3.68, Avicel PH-200 58.99, Me cellulose 0.07, and silica 0.81 mg.

L43 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2005 ACS on STN

1994:686615 CAPLUS ACCESSION NUMBER:

121:286615 DOCUMENT NUMBER:

TITLE: Pharmaceutical compositions containing pregnane

derivatives for the treatment of tumors.

INVENTOR(S): Deckers, Godefridus Hermanus Johanna; Kloosterboer,

Helenius Jan

PATENT ASSIGNEE(S): Akazo Nobel N.V., Neth.

Eur. Pat. Appl., 9 pp. SOURCE:

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 613687 EP 613687	A1 B1	19940907 19990602	EP 1994-200523	19940302
R: AT, BE, CH,	DE, DK	, ES, FR, GB	GR, IE, IT, LI, LU	J, MC, NL, PT, SE
CA 2116829	AA	19940906	CA 1994-2116829	19940302
CA 2116829	С	20040914		
AU 9457542	A1	19940908	AU 1994-57542	19940302
AU 671706	B2	19960905		
ZA 9401464	Α	19940927	ZA 1994-1464	19940302
AT 180669	E	19990615	AT 1994-200523	19940302
ES 2134313	Т3	19991001	ES 1994-200523	19940302
NO 9400777	A	19940906	NO 1994-777	19940304
JP 07002673	A2	19950106	JP 1994-34540	19940304
JP 3391539	B2	20030331		
US 5512556	A	19960430	US 1994-358639	19941219
PRIORITY APPLN. INFO.:			EP 1993-200631	A 19930305
			US 1994-206994	B1 19940307

OTHER SOURCE(S): MARPAT 121:286615

Entered STN: 10 Dec 1994 ED

Pharmaceutical compns. containing pregnane derivs. (Markush structure given) are useful for prevention or treatment of tumors. A twice daily dose of ${\bf 1}$ AB mg/kg $(7\alpha, 17\alpha)-17-hydroxy-7-methyl-19-nor-17-pregn-5(10)-en-20-mg/kg$ yn-3-one (I) inhibited the tumor growth in rats up to .apprx.70%. A tablet contained I 2.5, starch 10, ascorbyl palmitate 0.2, Mg stearate 0.5, and lactose q.s. to 100 mg.

L43 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1991:214521 CAPLUS

DOCUMENT NUMBER:

114:214521

TITLE:

Pharmaceutical composition containing the pure

crystalline forms of $(7\alpha, 17\alpha)-17-hydroxy-$ 7-methyl-19-nor-17-pregn-5(10)-en-20-yn-3-oneSas, G. A. J. M. T.; Van Doornum, E. M.

INVENTOR(S): AKZO N. V., Neth.

PATENT ASSIGNEE(S):

Eur. Pat. Appl., 13 pp. SOURCE:

CODEN: EPXXDW

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

1	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
		A1 B1	19900926 19931215	EP 1990-200574	19900312
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2	ZA 9001674	Α	19901228	ZA 1990-1674	19900305
(CA 2011452	C	19990525	CA 1990-2011452	19900305
1	AT 98490	E	19940115	AT 1990-200574	19900312
I	ES 2062292	Т3	19941216	ES 1990-200574	19900312
Ç	JP 03047195	A2	19910228	JP 1990-63921	19900314
Ç	JP 3139709	B2	20010305		
Ç	JP 2001139473	A2	20010522	JP 2000-316716	19900314
ţ	JS 5037817	Α	19910806	US 1990-494089	19900315
I	AU 9051394	A1	19900920	AU 1990-51394	19900316
I	AU 625083	B2	19920702		
I	FI 94865	В	19950731	FI 1990-1328	19900316
I	FI 94865	С	19951110		
F	KR 183401	B1	19990501	KR 1990-3574	19900317
PRIOR	TY APPLN. INFO.:			NL 1989-673	A 19890318
				EP 1990-200574	A 19900312
				JP 1990-63921	A3 19900314
DD .		1001			

ED Entered STN: 31 May 1991

AB Pure monoclinic or triclinic $(7\alpha,17\alpha)-17-hydroxy-7-methyl-19-nor-17-pregn-5(10)-en-20-yn-3-one (I)$ are more suitable for pharmaceutical compns. than the polymorphous I. A solution of 1 kg I in 23 L acetone containing

6 mL pyridine, was poured, under N, in 25 L water, followed by cooling to 0-5°, to crystallize monoclinic I (crystalline structure and x-ray diffraction spectrum given). Tablets comprised monoclinic I 2.5, starch 10, ascorbyl palmitate 0.2, Mg stearate 0.5, and lactose to 100 mg/tablet. I forms are useful for treating menopausal disorders, for immunomodulation, and for treating osteoporosis (no data).

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L16

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1 S E4

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          61648 S LACTOSE OR DISACCHARIDE?
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              3 S TIBOLONE? AND STARCH?
L11
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              3 S E3, E5, E7
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             96 S E3-E9, E17-E18
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                E ZWINKELS J/AU
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L35
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              8 S L26 AND L30
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L37
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                E GRANULATED STARCH/CN
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CA SUBSCRIBER PRICE
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FILE 'REGISTRY' ENTERED AT 12:43:55 ON 25 MAR 2005
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
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STRUCTURE FILE UPDATES: 24 MAR 2005 HIGHEST RN 847222-24-6 DICTIONARY FILE UPDATES: 24 MAR 2005 HIGHEST RN 847222-24-6

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 18, 2005

Please note that search-term pricing does apply when conducting SmartSELECT searches.

************ * The CA roles and document type information have been removed from * * the IDE default display format and the ED field has been added, * effective March 20, 2005. A new display format, IDERL, is now * available and contains the CA role and document type information. * ******************

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

=> e	lactose/cn	
E1	1	LACTOSAZONE/CN
E2	1	LACTOSCATONE/CN
E3	1>	LACTOSE/CN
E4	1	LACTOSE 100M/CN
E5	1	LACTOSE 2',6'-DIPALMITATE/CN
E6	1	LACTOSE 2'-PALMITATE/CN
E7	1	LACTOSE 2,4°DINITROPHENYLHYDRAZONE/CN
E8	1	LACTOSE 3'-PHOSPHATE/CN
E9	1	LACTOSE 4'-PHOSPHATE/CN
E10	1	LACTOSE 4-NITROPHENYLHYDRAZONE/CN
E11	1	LACTOSE 6'-PALMITATE/CN
E12	1	LACTOSE 6'-PHOSPHATE/CN
=> s	e3	
1.44	1 LAC	TOSE/CN

=> file caplus COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 5.03 421.53 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE 0.00 -14.60

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FILE COVERS 1907 - 25 Mar 2005 VOL 142 ISS 14 FILE LAST UPDATED: 24 Mar 2005 (20050324/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 144 L45 23757 L44 => d his

(FILE 'HOME' ENTERED AT 12:12:09 ON 25 MAR 2005)

FILE 'REGISTRY' ENTERED AT 12:12:34 ON 25 MAR 2005 E TIBOLONE/CN

L1 2 S E2-E3

FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005 386 S L1

L5 147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE L6 348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP L7 78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC

L8 61648 S LACTOSE OR DISACCHARIDE?
L9 5 S L4 AND L5 AND (L6 OR L7)
L10 3 S TIBOLONE? AND STARCH?

L11 0 S L10 NOT L9
L12 7 S L4 AND L8
L13 2 S L12 NOT L9
E HAAN P/AU
L14 3 S E3,E5,E7
E DE HAAN P/AU

L15 96 S E3-E9, E17-E18 E LAMBREGTS T/AU

L16 1 S E4
E HURK T/AU
E VD HURK T/AU
E VAN DEN HURK T/AU

L19 1 S E4 E ZWINKELS J/AU L20 3 S E3, E8 L21 99 S L14 OR L15

L22 198 S L21 OR L16 OR L17 OR L18 OR L19 OR L20

L23 4 S L22 AND TIBOLONE?

FILE 'MEDLINE, BIOSIS, CAPLUS, EMBASE, WPIDS' ENTERED AT 12:29:23 ON 25 MAR 2005

L24 8818 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR BOLTIN? OR "ORG OD14" L25 2399 S L24 NOT BOLTIN?

L26 2401 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR "ORG OD14" OR "ORG OD14" L27 276310 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE

L28 693483 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP

```
L29
         176163 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L30
         139122 S LACTOSE OR DISACCHARIDE?
L31
              3 S L26 AND L27 AND (L28 OR L29)
L32
              3 DUP REM L31 (0 DUPLICATES REMOVED)
L33
              6 S TIBOLONE? AND STARCH?
L34
              5 DUP REM L33 (1 DUPLICATE REMOVED)
L35
              2 S L34 NOT L32
L36
              8 S L26 AND L30
L37
              5 DUP REM L36 (3 DUPLICATES REMOVED)
L38
              2 S L37 NOT L34
L39
              5 S "BOLTIN"
L40
              0 S L39 AND STARCH?
     FILE 'STNGUIDE' ENTERED AT 12:38:54 ON 25 MAR 2005
     FILE 'REGISTRY' ENTERED AT 12:40:55 ON 25 MAR 2005
                E POTATO STARCH/CN
L41
              1 S E3
                E CORN STARCH/CN
L42
              1 S E3
                E WHEAT STARCH/CN
                E MODIFIED STARCH/CN
                E AGGLOMERATED STARCH/CN
                E GRANULATED STARCH/CN
     FILE 'CAPLUS' ENTERED AT 12:42:18 ON 25 MAR 2005
L43
              6 S L4 AND (L5 OR L41 OR L42)
     FILE 'REGISTRY' ENTERED AT 12:43:55 ON 25 MAR 2005
                E LACTOSE/CN
L44
              1 S E3
     FILE 'CAPLUS' ENTERED AT 12:44:08 ON 25 MAR 2005
L45
          23757 S L44
=> s 14 and 145
L46
            0 L4 AND L45
=> s tibolone and starch?
           358 TIBOLONE
             1 TIBOLONES
           358 TIBOLONE
                 (TIBOLONE OR TIBOLONES)
        147702 STARCH?
L47
             3 TIBOLONE AND STARCH?
=> s 147 not 143
L48
             0 L47 NOT L43
=> d his
     (FILE 'HOME' ENTERED AT 12:12:09 ON 25 MAR 2005)
     FILE 'REGISTRY' ENTERED AT 12:12:34 ON 25 MAR 2005
                E TIBOLONE/CN
L1
              2 S E2-E3
     FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005
L2
            386 S L1
                E TIBOLONE/BI
L3
            359 S E2-E4
                E TIBOLONE/CT
            430 S L2 OR L3
L5
         147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
```

```
348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
L7
          78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L8
          61648 S LACTOSE OR DISACCHARIDE?
              5 S L4 AND L5 AND (L6 OR L7)
L9
              3 S TIBOLONE? AND STARCH?
L10
              0 S L10 NOT L9
L11
L12.
              7 S L4 AND L8
L13
              2 S L12 NOT L9
                E HAAN P/AU
              3 S E3, E5, E7
L14
                E DE HAAN P/AU
             96 S E3-E9, E17-E18
L15
                E LAMBREGTS T/AU
L16
              1 S E4
                E HURK T/AU
                E VD HURK T/AU
                E VAN DEN HURK T/AU
              1 S E4
L17
                E MORITA R/AU
             97 S E3-E6, E23
L18
                E ROVERS A/AU
              1 S E4
1.19
                E ZWINKELS J/AU
              3 S E3, E8
L20
L21
             99 S L14 OR L15
            198 S L21 OR L16 OR L17 OR L18 OR L19 OR L20
L22
L23
              4 S L22 AND TIBOLONE?
     FILE 'MEDLINE, BIOSIS, CAPLUS, EMBASE, WPIDS' ENTERED AT 12:29:23 ON 25
     MAR 2005
L24
           8818 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR BOLTIN? OR "ORG OD14"
           2399 S L24 NOT BOLTIN?
L25
           2401 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR "ORG OD14" OR "ORG OD14"
L26
         276310 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L27
         693483 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
L28
L29
         176163 S (ASCORBYL PALMITATE?) OR "6-0-PALMITOYLASCORBIC ACID" OR "ASC
         139122 S LACTOSE OR DISACCHARIDE?
L30
L31
              3 S L26 AND L27 AND (L28 OR L29)
              3 DUP REM L31 (0 DUPLICATES REMOVED)
L32
L33
             .6 S TIBOLONE? AND STARCH?
              5 DUP REM L33 (1 DUPLICATE REMOVED)
L34
L35
              2 S L34 NOT L32
L36
              8 S L26 AND L30
L37
              5 DUP REM L36 (3 DUPLICATES REMOVED)
L38
              2 S L37 NOT L34
              5 S "BOLTIN"
L39
L40
              0 S L39 AND STARCH?
     FILE 'STNGUIDE' ENTERED AT 12:38:54 ON 25 MAR 2005
     FILE 'REGISTRY' ENTERED AT 12:40:55 ON 25 MAR 2005
                E POTATO STARCH/CN
L41
              1 S E3
                E CORN STARCH/CN
              1 S E3
L42
                E WHEAT STARCH/CN
                E MODIFIED STARCH/CN
                E AGGLOMERATED STARCH/CN
                E GRANULATED STARCH/CN
     FILE 'CAPLUS' ENTERED AT 12:42:18 ON 25 MAR 2005
L43
              6 S L4 AND (L5 OR L41 OR L42)
```

FILE 'REGISTRY' ENTERED AT 12:43:55 ON 25 MAR 2005

L17

1 S E3

```
L44
     FILE 'CAPLUS' ENTERED AT 12:44:08 ON 25 MAR 2005
          23757 S L44
             0 S L4 AND L45
L46
L47
              3 S TIBOLONE AND STARCH?
L48
             0 S L47 NOT L43
=> save
ENTER L#, L# RANGE, ALL, OR (END):all
ENTER NAME OR (END):110754733/1
L# LIST L1-L48 HAS BEEN SAVED AS 'L10754733/L'
=> d cost
COST IN U.S. DOLLARS
                                                 SINCE FILE
                                                                TOTAL
                                                     ENTRY
                                                               SESSION
CONNECT CHARGES
                                                       0.78
                                                               45.71
NETWORK CHARGES
                                                                 3.42
                                                       0.12
SEARCH CHARGES
                                                       3.78
                                                                317.84
DISPLAY CHARGES
                                                       0.00
                                                    ____
                                                               _____
FULL ESTIMATED COST
                                                       4.68
                                                               426.21
                                                               TOTAL
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
                                                 SINCE FILE
                                                     ENTRY
                                                               SESSION
CA SUBSCRIBER PRICE
                                                       0.00
                                                              -14.60
IN FILE 'CAPLUS' AT 12:45:05 ON 25 MAR 2005
=> d his
     (FILE 'HOME' ENTERED AT 12:12:09 ON 25 MAR 2005)
     FILE 'REGISTRY' ENTERED AT 12:12:34 ON 25 MAR 2005
               E TIBOLONE/CN
              2 S E2-E3
L1
     FILE 'CAPLUS' ENTERED AT 12:12:50 ON 25 MAR 2005
            386 S L1
L2
               E TIBOLONE/BI
            359 S E2-E4
L3
               E TIBOLONE/CT
            430 S L2 OR L3
         147787 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L5
         348947 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
L6
          78296 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L7
          61648 S LACTOSE OR DISACCHARIDE?
L8
              5 S L4 AND L5 AND (L6 OR L7)
L9
              3 S TIBOLONE? AND STARCH?
L10
              0 S L10 NOT L9
L11
L12
              7 S L4 AND L8
              2 S L12 NOT L9
L13
                E HAAN P/AU
              3 S E3, E5, E7
L14
                E DE HAAN P/AU
L15
             96 S E3-E9, E17-E18
               E LAMBREGTS T/AU
              1 S E4
L16
                E HURK T/AU
                E VD HURK T/AU
```

E VAN DEN HURK T/AU

1 S E4

```
E MORITA R/AU
             97 S E3-E6, E23
L18
                E ROVERS A/AU
L19
              1 S E4
                E ZWINKELS J/AU
L20
              3 S E3, E8
L21
             99 S L14 OR L15
L22
            198 S L21 OR L16 OR L17 OR L18 OR L19 OR L20
L23
              4 S L22 AND TIBOLONE?
     FILE 'MEDLINE, BIOSIS, CAPLUS, EMBASE, WPIDS' ENTERED AT 12:29:23 ON 25
     MAR 2005
           8818 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR BOLTIN? OR "ORG OD14"
L24
           2399 S L24 NOT BOLTIN?
L25
           2401 S TIBOLONE? OR LIVIAL? OR LIVIELLA? OR "ORG OD14" OR "ORG OD14"
L26
         276310 S STARCH? OR (POTATO STARCH?) OR (CORN STARCH?) OR (AMYLOMAIZE
L27
L28
         693483 S STABILIZER? OR (STABILIZING (W) (AGENT? OR COMPONENT? OR COMP
        176163 S (ASCORBYL PALMITATE?) OR "6-O-PALMITOYLASCORBIC ACID" OR "ASC
L29
         139122 S LACTOSE OR DISACCHARIDE?
L30
L31
              3 S L26 AND L27 AND (L28 OR L29)
L32
              3 DUP REM L31 (0 DUPLICATES REMOVED)
              6 S TIBOLONE? AND STARCH?
L33
L34
              5 DUP REM L33 (1 DUPLICATE REMOVED)
L35
             2 S L34 NOT L32
L36
             8 S L26 AND L30
L37
             5 DUP REM L36 (3 DUPLICATES REMOVED)
L38
             2 S L37 NOT L34
L39
              5 S "BOLTIN"
L40
              0 S L39 AND STARCH?
     FILE 'STNGUIDE' ENTERED AT 12:38:54 ON 25 MAR 2005
     FILE 'REGISTRY' ENTERED AT 12:40:55 ON 25 MAR 2005
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L41
              1 S E3
                E CORN STARCH/CN
L42
              1 S E3
                E WHEAT STARCH/CN
                E MODIFIED STARCH/CN
                E AGGLOMERATED STARCH/CN
                E GRANULATED STARCH/CN
     FILE 'CAPLUS' ENTERED AT 12:42:18 ON 25 MAR 2005
L43
              6 S L4 AND (L5 OR L41 OR L42)
     FILE 'REGISTRY' ENTERED AT 12:43:55 ON 25 MAR 2005
                E LACTOSE/CN
              1 S E3
L44
     FILE 'CAPLUS' ENTERED AT 12:44:08 ON 25 MAR 2005
L45
          23757 S L44
              0 S L4 AND L45
L46
              3 S TIBOLONE AND STARCH?
L47
L48
              0 S L47 NOT L43
                SAVE ALL L10754733/L
```

DERWENT

_						
L9	2814	11 and 17 and (15 or 16)	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2005/03/25 09:07
L10	187	11 and 17 and (15 or 16).ti,ab,clm.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2005/03/25 09:07
L11	234	tibolone or livial or liviella or (org adj OD14)	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/25 09:12
L12	116	I11 and I7 and (I5 or I6)	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/25 09:12
L13	20	I11 and I7 and (I5 or I6).ti,ab,clm.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/25 09:31
L14	1173	I1 not I11	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/25 09:31
L15	2	(I1 not I11).ti,ab,clm.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/25 09:31
L16	130354	lactose or disaccharide	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2005/03/25 09:49
L17	96	I12 not I13	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2005/03/25 10:59
L18	101	111 and 17 and (15 or 16) and 116	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2005/03/25 10:59

L19	0	l18 not l12	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2005/03/25 10:59
L20	69	tibolone and starch and (ascorb\$)	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2005/03/25 11:00
L21	0	I20 not I12	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2005/03/25 11:00
L22	13	("4473564" "4172132" "3423507" "4185101" "4159326").pn.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2005/03/25 11:01
S1	3	("6514958" "6399594").pn.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/25 08:58
S2	2	("6156343").pn.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR .	OFF	2005/03/24 11:27
S3	2	("5916593").pn.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 11:12
S4	20	("4708874" "5178870" "5292520" "5441747" "5382434" "5662936" "6187339" "5916593" "5709881" "6063403").pn.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 11:12
S5	2	("4701450").pn.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 11:32
S6	0	ep-0389035-\$.did.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 11:32

S7	2	ep-389035-\$.did.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 11:37
S8	2	ep-613687-\$.did.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 11:39
S9	2	ep-707848-\$.did.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 11:43
S10	2	ep-503521-\$.did.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 11:53
S11	2	wo-9506461-\$.did.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 15:24
S12	228	tibolone	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 15:24
S13	161	S12 and starch	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 15:24
S14	9	(tibolone and starch and (antioxidant or chelat\$)).ti,ab,clm.	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	OFF	2005/03/24 15:25